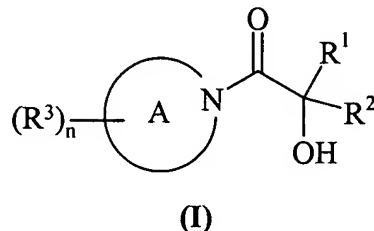


**IN THE CLAIMS:**

Claim 1 (currently amended): A compound of formula (I):



wherein:

**Ring A** is piperazinyl ~~optionally~~ substituted on nitrogen by  $R^4$ -D-;

$R^1$  and  $R^2$  are independently  $C_k$ alkyl optionally substituted by 1 to  $2k+1$  atoms selected from fluoro and chloro wherein  $k$  is 1-3;

or  $R^1$  and  $R^2$  together with the carbon atom to which they are attached, form a  $C_m$ cycloalkyl ring optionally substituted by 1 to  $2m-2$  fluorine atoms wherein  $m$  is 3-5;

$R^3$  is a substituent on carbon and is halo, hydroxy, cyano, formyl, amino, nitro, carboxy, carbamoyl, ureido, thiol, sulphamoyl or  $R^5$ -E-;

$R^4$  is  $C_{1-6}$ alkyl, phenyl or a heterocyclic group, wherein in  $R^4$  any  $C_{1-6}$ alkyl, phenyl or heterocyclic groups (on a ring carbon) may be optionally substituted by one or more  $R^6$  and if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from  $R^8$ ;

**D** is  ~~$C(O)$ -,  $-N(R^9)C(O)$ -,  $-S(O)_2$ - or  $-NS(O)_2$ -,  $-OC(O)$ -~~ or **D** is a direct bond;

$R^5$  is  $C_{1-6}$ alkyl,  $C_{2-6}$ alkenyl,  $C_{2-6}$ alkynyl,  $C_{3-6}$ cycloalkyl, phenyl, naphthyl or a heterocyclic group, wherein in  $R^5$  any  $C_{1-6}$ alkyl,  $C_{2-6}$ alkenyl,  $C_{2-6}$ alkynyl,  $C_{3-6}$ cycloalkyl, phenyl, naphthyl or heterocyclic groups (on a ring carbon) may be optionally substituted by one or more  $R^6$  and if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from  $R^8$ ;

**E** is -O-,  $-N(R^9)$ -,  $-C(O)$ -,  $-N(R^9)C(O)$ -,  $-C(O)N(R^9)$ -,  $-S(O)_a$ - wherein  $a$  is 0-2,  $-OC(O)$ -,  $-C(O)O$ -,  $-N(R^9)C(O)O$ -,  $-OC(O)N(R^9)$ -,  $-C(S)N(R^9)$ -,  $-N(R^9)C(S)$ -,  $-SO_2N(R^9)$ -,  $-N(R^9)SO_2$ -,  $-N(R^9)C(O)N(R^9)$ -,  $-N(R^9)C(S)N(R^9)$ -,  $-SO_2NHC(O)$ -,  $-SO_2N(R^9)C(O)$ -,  $-C(O)NHSO_2$ - or **E** is a direct bond;

**R<sup>6</sup>** is trifluoromethyl, C<sub>1-6</sub>alkyl, halo, hydroxy, trifluoromethoxy, cyano, C<sub>1-6</sub>alkoxy, formyl, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkanoyloxy, amino, *N*-(C<sub>1-6</sub>alkyl)amino, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>amino, C<sub>1-6</sub>alkanoylamino, C<sub>1-6</sub>alkanoyl(*N*-C<sub>1-6</sub>alkyl)amino, nitro, carboxy, carbamoyl, C<sub>1-6</sub>alkoxycarbonyl, thiol, C<sub>1-6</sub>alkylsulphanyl, C<sub>1-6</sub>alkylsulphinyl, C<sub>1-6</sub>alkylsulphonyl, C<sub>1-6</sub>alkylsulphonylamino, sulphamoyl, *N*-(C<sub>1-6</sub>alkyl)aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, ureido, *N'*-(C<sub>1-6</sub>alkyl)ureido or *N'*-(C<sub>1-6</sub>alkyl)<sub>2</sub>ureido, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl or C<sub>3-6</sub>cycloalkyl, naphthyl, phenyl or a heterocyclic group wherein in **R<sup>6</sup>** any C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, C<sub>3-6</sub>cycloalkyl, naphthyl, phenyl or heterocyclic groups (on a ring carbon) may be optionally substituted by one or more **R<sup>7</sup>** and if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from **R<sup>8</sup>**;

**R<sup>7</sup>** is trifluoromethyl, cyano, C<sub>1-6</sub>alkyl, halo, hydroxy, trifluoromethoxy, C<sub>1-6</sub>alkoxy, formyl, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkanoyloxy, amino, *N*-(C<sub>1-6</sub>alkyl)amino, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>amino, C<sub>1-6</sub>alkanoylamino, C<sub>1-6</sub>alkanoyl(*N*-C<sub>1-6</sub>alkyl)amino, nitro, carboxy, carbamoyl, C<sub>1-6</sub>alkoxycarbonyl, thiol, C<sub>1-6</sub>alkylsulphanyl, C<sub>1-6</sub>alkylsulphinyl, C<sub>1-6</sub>alkylsulphonyl, C<sub>1-6</sub>alkylsulphonylamino, sulphamoyl, *N*-(C<sub>1-6</sub>alkyl)aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, C<sub>3-6</sub>cycloalkyl or a heterocyclic group (optionally substituted by one or more **R<sup>11</sup>**), and wherein in **R<sup>7</sup>** any C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl or C<sub>3-6</sub>cycloalkyl groups may be optionally substituted by one or more groups selected from **R<sup>12</sup>**;

**R<sup>8</sup>** is C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkylsulphonyl, C<sub>1-6</sub>alkoxycarbonyl, carbamoyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N,N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, benzoyl, (heterocyclic group)carbonyl, phenylsulphonyl, (heterocyclic group)sulphonyl, phenyl or a carbon linked heterocyclic group, and wherein in **R<sup>8</sup>** any C<sub>1-6</sub>alkyl, phenyl or heterocyclic group (on a ring carbon) may be optionally substituted by one or more **R<sup>6</sup>**, and if a heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from **R<sup>11</sup>**;

wherein for **R<sup>4</sup>**, **R<sup>5</sup>**, **R<sup>6</sup>**, **R<sup>7</sup>** and **R<sup>8</sup>**, a heterocyclic group is selected from morpholino, piperidyl, pyridyl, pyranyl, pyrrolyl, isothiazolyl, indolyl, quinolyl, thienyl,

1,3-benzodioxolyl, thiadiazolyl, piperazinyl, thiazolidinyl, pyrrolidinyl, thiomorpholino, pyrrolinyl, homopiperazinyl, tetrahydropyranyl, imidazolyl, pyrimidyl, pyrazinyl, pyridazinyl, isoxazolyl, 4-pyridone, 1-isoquinolone, 2-pyrrolidone, 4-thiazolidone, pyridine-*N*-oxide, quinoline-*N*-oxide and combinations thereof;

**R<sup>9</sup>** is hydrogen or C<sub>1-6</sub>alkyl optionally substituted by one or more R<sup>10</sup> with the proviso that R<sup>10</sup> is not a substituent on the carbon attached to a nitrogen atom;

**R<sup>10</sup>** is halo, hydroxy, amino, cyano, nitro, trifluoromethyl, trifluoromethoxy, C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, *N*-(C<sub>1-6</sub>alkyl)amino, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>amino, C<sub>1-6</sub>alkanoylamino, C<sub>1-6</sub>alkanoyl(*N*-C<sub>1-6</sub>alkyl)amino, C<sub>1-6</sub>alkylsulphonylamino, C<sub>1-6</sub>alkylsulphonyl(*N*-C<sub>1-6</sub>alkyl)amino, thiol, C<sub>1-6</sub>alkylsulphanyl, C<sub>1-6</sub>alkylsulphinyl, C<sub>1-6</sub>alkylsulphonyl, sulphamoyl, *N*-(C<sub>1-6</sub>alkyl)aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>aminosulphonyl, carboxy, carbamoyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-6</sub>alkoxycarbonyl, C<sub>1-6</sub>alkanoyl or formyl;

**R<sup>11</sup>** is C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkylsulphonyl, C<sub>1-6</sub>alkoxycarbonyl, carbamoyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N,N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-6</sub>alkoxyC<sub>1-6</sub>alkanoyl, phenylC<sub>1-6</sub>alkyl, benzoyl, phenylC<sub>1-6</sub>alkanoyl, phenylC<sub>1-6</sub>alkoxycarbonyl or phenylsulphonyl and wherein in R<sup>11</sup> any C<sub>1-6</sub>alkyl group can be optionally substituted by one or more R<sup>13</sup>;

**R<sup>12</sup>** is halo, hydroxy, *N*-methylpiperazinyl, *N*-acetylpiperazinyl, morpholino, piperidino, cyano, amino, *N,N*-dimethylamino, acetamido, carbamoyl, carboxy, methanesulphonyl or sulphamoyl;

**R<sup>13</sup>** is halo, hydroxy, cyano, amino, *N,N*-dimethylamino, acetamido, carbamoyl, carboxy, methanesulphonyl or sulphamoyl;

**n** is 0-5; wherein the values of R<sup>3</sup> may be the same or different;

or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof;

~~with the proviso that if R<sup>1</sup> is methyl, R<sup>2</sup> is trifluoromethyl and Ring A is piperazin-1-yl then (R<sup>3</sup>)<sub>n</sub> is not i) 4-cyanobenzoyl, ii) 2-methyl-4-benzyloxycarbonyl, iii) 2-methyl, iv) 2-methyl-4-cyanobenzoyl, v) 2,5-dimethyl-4-benzyl, vi) 2,5-dimethyl or vii) 2,5-dimethyl-4-cyanobenzoyl.~~

Claim 2 (original): A compound of formula (I) according to claim 1 wherein one of R<sup>1</sup> and R<sup>2</sup> is methyl and the other is trifluoromethyl;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 3 (cancelled).

Claim 4 (previously presented): A compound of formula (I) according to claim 1 wherein R<sup>3</sup> is a substituent on carbon and is selected from amino, methyl, 4-mesylphenylsulphonyl, 4-methylthiophenylthio, 4-fluorobenzoyl and 4-cyanobenzoylamino;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 5 (previously presented): A compound of formula (I) according to claim 1 wherein R<sup>4</sup> is C<sub>1-4</sub>alkyl, phenyl {optionally substituted with one or more *t*-butyl, isopropyl, nitro, halo, *N,N*-dimethylcarbamoyl, *N,N*-dimethylamino, 2-hydroxyethylamino, cyano, acetyl, methoxy or carboxy} or thienyl;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 6 (currently amended): A compound of formula (I) according to claim 1 wherein D is ~~-SO<sub>2</sub>-~~or ~~C(O)-~~;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 7 (previously presented): A compound of formula (I) according to claim 1 wherein n is 0 - 3;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 8 (currently amended): A compound of formula (I) according to claim 1,  
selected from:

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-carboxyphenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-dimethylcarbamoylphenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-fluorophenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

(R)-{(2S,5R)-2-methyl-5-methyl-4-[4-(2-hydroxyethylamino)phenylsulphonyl]-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine};

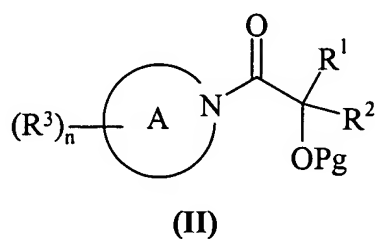
(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-cyanophenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine]; and

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-methoxyphenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

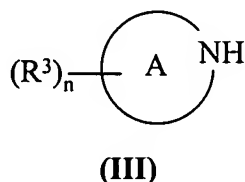
Claim 9 (currently amended): A process for preparing a compound of formula (I) as described in claim 1, or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, which process (in which variable groups are as defined in claim 1 for formula (I) unless otherwise stated) comprises of:

(a) deprotecting a protected compound of formula (II):

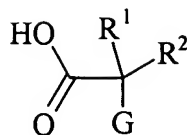


where Pg is an alcohol protecting group;

(b) coupling an amine of formula (III):



with an acid of formula (IV):



(IV)

wherein G is a hydroxyl group;

(c) coupling an amine of formula (III) with an activated acid derivative of formula (IV)

wherein G is a hydroxyl group which may be protected as an ester or ether;

and thereafter if necessary:

- i) converting a compound of the formula (I) into another compound of the formula (I);
- ii) removing any protecting groups; or
- iii) forming a pharmaceutically acceptable salt or *in vivo* hydrolysable ester.

Claim 10 (previously presented): A pharmaceutical composition which comprises a compound of formula (I) according to any one of claims 1-2 and 4-8, or a pharmaceutically acceptable salt or *in vivo* hydrolysable ester thereof in association with a pharmaceutically-acceptable diluent or carrier.

Claims 11-15 (cancelled).

Claim 16 (new): A method for the treatment of diabetes mellitus, said method comprising administering to a warm-blooded animal in need thereof a diabetes mellitus effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 17 (new): A method for the treatment of peripheral vascular disease, said method comprising administering to a warm-blooded animal in need thereof a peripheral vascular disease effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 18 (new): A method for the treatment of ischaemia, said method comprising administering to a warm-blooded animal in need thereof an ischaemia effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 19 (new): A method for the treatment of hyperlipidaemia, said method comprising administering to a warm-blooded animal in need thereof a hyperlipidaemia effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 20 (new): A method for the treatment of Alzheimers disease, said method comprising administering to a warm-blooded animal in need thereof an Alzheimers disease effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 21 (new): A method for the treatment of atherosclerosis, said method comprising administering to a warm-blooded animal in need thereof an atherosclerosis effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.